

论文

速激肽NK-1受体拮抗剂SR-140333对抗原引起致敏大鼠气道高反应性的影响

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摘要:

为观察速激肽NK-1受体拮抗剂SR-140333对抗原攻击引起的致敏大鼠气道高反应性的影响,测定了致敏大鼠在抗原攻击前后的基础呼吸频率,对MCh的反应性及支气管-肺泡灌洗液中的白细胞数量。实验结果显示,致敏大鼠吸入OA后6h基础呼吸频率增加,并显著增加乙酰甲胆碱(MCh)的反应性、MCh的-logPC₃₀值和支气管-肺泡灌洗液中的白细胞数量。ip速激肽NK-1受体拮抗剂SR-140333(0.1mg·kg⁻¹)或地塞米松(0.5mg·kg⁻¹),可明显抑制上述反应,小剂量SR-140333(0.01mg·kg⁻¹)仅有部分抑制作用。结果提示抗原攻击可引起致敏大鼠气道高反应性和气道炎症,速激肽NK-1受体拮抗剂可抑制这些反应。

关键词: 气道高反应性 NK-1受体拮抗剂SR-140333 速激肽受体拮抗剂 地塞米松 气道炎症

EFFECT OF SR-140333, A TACHYKININ NK-1 ANTAGONIST, ON ANTIGEN-INDUCED AIRWAY HYPERRESPONSIVENESS IN SENSITIZED RATS

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Abstract:

In the present study, the effects of SR-140333, { (S)-1-(2-[3,4-dichlorophenyl]-1-(3-isopropoxyphenylacetyl)piperidin-3yl)ethyl)-4-phenyl-1-azoniabicyclooctane-chloride } , a nonpeptide antagonist for tachykinin NK-1 receptor, on the antigen-induced airway response to methacholine (MCh) aerosol and airway inflammation in sensitized SD rats were investigated. The baseline respiratory frequencies, tachypnea response to methacholine(MCh), the-log PC₃₀ values of MCh and the leukocyte counts in bronchoalveolar lavage significantly increased after inhalation of 1% oval albumin(OA) aerosol. SR-140333 (152 nmol·kg⁻¹, ip) or dexamethasone(368 nmol·kg⁻¹, ip), bid×3 d inhibited these responses. SR-140333 at a low dose of 0.01 mg·kg⁻¹ showed an incomplete inhibition. From these results, we conclude that antigen challenge causes airway hyperresponsiveness and airway inflammation and that tachykinin NK-1 receptor antagonist inhibits these responses.

Keywords: SR-140333 Tachykinin NK-1 receptor antagonist Dexamethasone Airway inflammation Airway hyperresponsiveness

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